

MAR 0 7 7000 5

<110> McCarthy, Sean A.

<120> NOVEL HUMAN DICKKOPF-RELATED PROTEIN AND NUCLEIC ACID MOLECULES AND USES THEREFOR

<130> MNI-108CP2

<140> 09/263,022

<141> 1999-03-05

<150> 08/843,704

<151> 1997-04-16

<150> 08/842,898

<151> 1997-04-17

<150> 60/071,589

<151> 1998-01-15

<150> 09/009,802

<151> 1998-01-20

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<170> PatentIn Ver. 2.0

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<222> (38)..(1087)

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acc ctg ctg tgc ctg ctg gcg gcg gcg gtc.ccc acg gcc ccc gcg 103
Thr Leu Leu Cys Leu Leu Leu Ala Ala Ala Val Pro Thr Ala Pro Ala
10 15 20

ccc gct ccg acg gcg acc tcg gct cca gtc aag ccc ggc ccg gct ctc 151
Pro Ala Pro Thr Ala Thr Ser Ala Pro Val Lys Pro Gly Pro Ala Leu
25 30 35

agc tac ccg cag gag gac gcc acc ctc aat gag atg ttc cgc gag gtt 199 Ser Tyr Pro Gln Glu Glu Ala Thr Leu Asn Glu Met Phe Arg Glu Val 40 45 50

 \mathcal{N}^{1}

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				gaa Glu 75										295
				cct Pro										343
				aat Asn										391
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				ccc Pro 155										535
	_	_		tgc Cys			_	 _						583
				gac Asp										631
_				agc Ser				_	_	_		_	_	679
				tgc Cys								Pro		727
				ccc Pro 235										775
		_	_	ctc Leu						-				823
				tgt Cys										871
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agc ctg act gaa gag atg gcg ctg agg gag cct gcg gct gcc gcc gct 1063 Ser Leu Thr Glu Glu Met Ala Leu Arg Glu Pro Ala Ala Ala Ala Ala 330 335 340
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Lys Pro Gly Pro Ala Leu Ser Tyr Pro Gln Glu Glu Ala Thr Leu Asn 35 40 45

Glu Met Phe Arg Glu Val Glu Glu Leu Met Glu Asp Thr Gln His Lys 50 55 60

Leu Arg Ser Ala Val Glu Glu Met Glu Ala Glu Glu Ala Ala Ala Lys
65 70 75 80

Ala Ser Ser Glu Val Asn Leu Ala Asn Leu Pro Pro Ser Tyr His Asn 85 90 95

Glu Thr Asn Thr Asp Thr Asn Val Gly Asn Asn Thr Ile His Val His $100 \\ 0.5 \\ 105 \\ 110$

Arg Glu Ile His Lys Ile Thr Asn Asn Gln Thr Gly Gln Met Val Phe 115 120 125

Ser Glu Thr Val Ile Thr Ser Val Gly Asp Glu Gly Arg Arg Ser 130 135 140

His Glu Cys Ile Ile Asp Glu Asp Cys Gly Pro Ser Met Tyr Cys Gln 145 150 155 160

Phe Ala Ser Phe Gln Tyr Thr Cys Gln Pro Cys Arg Gly Gln Arg Met 165 170 175

Leu Cys Thr Arg Asp Ser Glu Cys Cys Gly Asp Gln Leu Cys Val Trp 180 185 190

Gly His Cys Thr Lys Met Ala Thr Arg Gly Ser Asn Gly Thr Ile Cys 195 200 205

Asp Asn Gln Arg Asp Cys Gln Pro Gly Leu Cys Cys Ala Phe Gln Arg 210 215 220

Gly Leu Leu Phe Pro Val Cys Thr Pro Leu Pro Val Glu Gly Glu Leu 225 230 235 240

Cys His Asp Pro Ala Ser Arg Leu Leu Asp Leu Ile Thr Trp Glu Leu 245 250 255

Glu Pro Asp Gly Ala Leu Asp Arg Cys Pro Cys Ala Ser Gly Leu Leu 260 265 270

Cys Gln Pro His Ser His Ser Leu Val Tyr Val Cys Lys Pro Thr Phe 275 280 285

Val Gly Ser Arg Asp Gln Asp Gly Glu Ile Leu Leu Pro Arg Glu Val 295 300 Pro Asp Glu Tyr Glu Val Gly Ser Phe Met Glu Glu Val Arg Gln Glu Leu Glu Asp Leu Glu Arg Ser Leu Thr Glu Glu Met Ala Leu Arg Glu 325 330 Pro Ala Ala Ala Ala Ala Leu Leu Gly Arg Glu Glu Ile 345 <210> 3 <211> 1050 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (1)..(1050) atg cag cgg ctt ggg gcc acc ctg ctg tgc ctg ctg gcg gcg gcg 48 Met Gln Arg Leu Gly Ala Thr Leu Leu Cys Leu Leu Leu Ala Ala Ala qtc ccc acq qcc ccc gcg ccc gct ccg acg gcg acc tcg gct cca gtc 96 Val Pro Thr Ala Pro Ala Pro Ala Pro Thr Ala Thr Ser Ala Pro Val aaq cee qqe eeq qet ete age tae eeq eaq gag gag gee ace ete aat 144 Lys Pro Gly Pro Ala Leu Ser Tyr Pro Gln Glu Glu Ala Thr Leu Asn gag atg ttc cgc gag gtt gag gaa ctg atg gag gac acg cag cac aaa .192Glu Met Phe Arg Glu Val Glu Glu Leu Met Glu Asp Thr Gln His Lys ttg cgc agc gcg gtg gaa gag atg gag gca gaa gaa gct gct gct aaa 240 Leu Arg Ser Ala Val Glu Glu Met Glu Ala Glu Glu Ala Ala Lys 70 gca tca tca gaa gtg aac ctg gca aac tta cct ccc agc tat cac aat 288 Ala Ser Ser Glu Val Asn Leu Ala Asn Leu Pro Pro Ser Tyr His Asn 90 85 gag acc aac aca gac acg aac gtt gga aat aat acc atc cat gtg cac 336 Glu Thr Asn Thr Asp Thr Asn Val Gly Asn Asn Thr Ile His Val His 105 100 384 cga gaa att cac aag ata acc aac aac cag act gga caa atg gtc ttt Arg Glu Ile His Lys Ile Thr Asn Asn Gln Thr Gly Gln Met Val Phe 120 125 432 tca gag aca gtt atc aca tct gtg gga gac gaa gaa ggc aga agg agc

Ser Glu Thr Val Ile Thr Ser Val Gly Asp Glu Glu Gly Arg Arg Ser

140

135

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_	_	_	tac Tyr	_	_		_	 	_	 -	528
			agt Ser					Leu			576
			atg Met								624
			tgc Cys								672
			gtg Val 230								720
			agc Ser								768
			ttg Leu								816
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Thr Arg Lys Phe Cys Leu Gln Pro Arg Asp Glu Lys Pro Phe Cys Ala 50 55 60

Thr Cys Arg Gly Leu Arg Arg Cys Gln Arg Asp Ala Met Cys Cys 65 70 75 80

Pro Gly Thr Leu Cys Val Asn Asp Val Cys Thr Thr Met Glu Asp Ala 85 90 95

Thr Pro Ile Leu Glu Arg Gln Leu Asp Glu Gln Asp Gly Thr His Ala 100 105 110

Glu Gly Thr Thr Gly His Pro Val Gln Glu Asn Gln Pro Lys Arg Lys 115 120 125

Pro Ser Ile Lys Lys Ser Gln Gly Arg Lys Gly Gln Glu Gly Glu Ser 130 135 140

Cys Leu Arg Thr Phe Asp Cys Gly Pro Gly Leu Cys Cys Ala Arg His 145 150 155 160

Phe Trp Thr Lys Ile Cys Lys Pro Val Leu Leu Glu Gly Gln Val Cys 165 170 175

Ser Arg Arg Gly His Lys Asp Thr Ala Gln Ala Pro Glu Ile Phe Gln 180 185 190

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cat His	ggg Gly	gcc Ala 35	cgg Arg	aag Lys	ggc Gly	tca Ser	cag Gln 40	tgc Cys	ctg Leu	tct Ser	gac Asp	acg Thr 45	gac Asp	tgc Cys	aat Asn	144
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aca Thr 65	tgt Cys	cgt Arg	ggg ggg	ttg Leu	cgg Arg 70	agg Arg	agg Arg	tgc Cys	cag Gln	cga Arg 75	gat Asp	gcc Ala	atg Met	tgc Cys	tgc Cys 80	240
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					cac His											384
					tca Ser											432
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cgt Arg	tgc Cys	gac Asp 195	tgt Cys	ggc Gly	cct Pro	gga Gly	cta Leu 200	ctg Leu	tgt Cys	cga Arg	agc Ser	caa Gln 205	ttg Leu	acc Thr	agc Ser	624
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170

165

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Leu Gly Gly Ala Ala Gly His Pro Gly Ser Ala Val Ser Ala Ala Pro Gly Ile Leu Tyr Pro Gly Gly Asn Lys Tyr Gln Thr Ile Asp Asn Tyr Gln Pro Tyr Pro Cys Ala Glu Asp Glu Glu Cys Gly Thr Asp Glu Tyr Cys Ala Ser Pro Thr Arg Gly Gly Asp Ala Gly Val Gln Ile Cys Leu Ala Cys Arg Lys Arg Lys Arg Cys Met Arg His Ala Met Cys Cys Pro Gly Asn Tyr Cys Lys Asn Gly Ile Cys Val Ser Ser Asp Gln Asn His Phe Arg Gly Glu Ile Glu Glu Thr Ile Thr Glu Ser Phe Gly Asn 150 Asp His Ser Thr Leu Asp Gly Tyr Ser Arg Arg Thr Thr Leu Ser Ser Lys Met Tyr His Thr Lys Gly Gln Glu Gly Ser Val Cys Leu Arg Ser 180 Ser Asp Cys Ala Ser Gly Leu Cys Cys Ala Arg His Phe Trp Ser Lys 200 Ile Cys Lys Pro Val Leu Lys Glu Gly Gln Val Cys Thr Lys His Arg 210 215 Arg Lys Gly Ser His Gly Leu Glu Ile Phe Gln Arg Cys Tyr Cys Gly 235 230 Glu Gly Leu Ser Cys Arg Ile Gln Lys Asp His His Gln Ala Ser Asn 250 245 Ser Ser Arg Leu His Thr Cys Gln Arg His 260 <210> 9 <211> 798 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (1)..(798) <400> 9 48 atg atg gct ctg ggc gca gcg gga gct acc cgg gtc ttt gtc gcg atg Met Met Ala Leu Gly Ala Ala Gly Ala Thr Arg Val Phe Val Ala Met 96 gta gcg gcg gct ctc ggc ggc cac cct ctg ctg gga gtg agc gcc acc Val Ala Ala Leu Gly Gly His Pro Leu Leu Gly Val Ser Ala Thr

20

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gga Gly 65	atc Ile	ctg Leu	tac Tyr	ccg Pro	ggc Gly 70	ggg Gly	aat Asn	aag Lys	tac Tyr	cag Gln 75	acċ Thr	att Ile	gac Asp	aac Asn	tac Tyr 80	240
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aga Arg 225	aaa Lys	ggc Gly	tct Ser	cat His	gga Gly 230	cta Leu	gaa Glu	ata Ile	ttc Phe	cag Gln 235	cgt Arg	tgt Cys	tac Tyr	tgt Cys	gga Gly 240	720
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cca Pro	gtt Val 50	act Thr	gaa Glu	agc Ser	atc Ile	tta Leu 55	acc Thr	cct Pro	cac His	atc Ile	ccg Pro 60	gct Ala	ctg Leu	gat Asp	ggt Gly	192
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ctg Leu 145	gaa Glu	att Ile	ttc Phe	cag Gln	cgt Arg 150	tgc Cys	gac Asp	tgt Cys	gcg Ala	aag Lys 155	ggc Gly	ctg Leu	tct Ser	tgc Cys	aaa. Lys 160	480
gta Val	tgg Trp	aaa Lys	gat Asp	gcc Ala 165	acc Thr	tac Tyr	tcc Ser	tcc Ser	aaa Lys 170	gcc Ala	aga Arg	ctc Leu	cat His	gtg Val 175	tgt Cys	528
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gtt	gtgt	att	taat	gcat [.]	ta t	agca	tggt	g ga	aaat	aagg	ttc	agat	gca	gaag	aatggc	637
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<213> Homo sapiens

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Asp Gly Met Cys Cys Pro Ser Thr Arg Cys Asn Asn Gly Ile Cys Ile 35 40 45

Pro Val Thr Glu Ser Ile Leu Thr Pro His Ile Pro Ala Leu Asp Gly 50 60

Thr Arg His Arg Asp Arg Asn His Gly His Tyr Ser Asn His Asp Leu 65 70 75 80

Gly Trp Gln Asn Leu Gly Arg Pro His Thr Lys Met Ser His Ile Lys 85 90 95

Gly His Glu Gly Asp Pro Cys Leu Arg Ser Ser Asp Cys Ile Glu Gly 100 105 110

Phe Cys Cys Ala Arg His Phe Trp Thr Lys Ile Cys Lys Pro Val Leu 115 120 125

His Gln Gly Glu Val Cys Thr Lys Gln Arg Lys Lys Gly Ser His Gly 130 135 140

Leu Glu Ile Phe Gln Arg Cys Asp Cys Ala Lys Gly Leu Ser Cys Lys 145 150 155 160

Val Trp Lys Asp Ala Thr Tyr Ser Ser Lys Ala Arg Leu His Val Cys 165 170 175

Gln Lys Ile

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gat Asp	ggc Gly	atg Met 35	tgc Cys	tgc Cys	ccc Pro	agt Ser	acc Thr 40	cgc Arg	tgc Cys	aat Asn	aat Asn	ggc Gly 45	atc Ile	tgt Cys	atc Ile	144
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cac His	cag Gln	ctg Leu 95	Gly	aac Asn	aac Asn	acc Thr	ctc Leu 100	tcc Ser	agc Ser	cac His	ctc Leu	cag Gln 105	atc Ile	gac Asp	aag Lys	398
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gca Ala 125	tcc Ser	att Ile	caa Gln	cca Pro	gcg Ala 130	gag Glu	ggg Gly	agc Ser	ttc Phe	gag Glu 135	ggt Gly	gat Asp	ttg Leu	aag Lys	gta Val 140	494
ccc Pro	agg Arg	atg Met	gag Glu	gag Glu 145	aag Lys	gag Glu	gcc Ala	ctg Leu	gta Val 150	ccc Pro	atc Ile	cag Gln	aag Lys	gcc Ala 155	acg Thr	542
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tcc Ser	càc His	tcc Ser	agg Arg	ctg Leu 225	tcc Ser	ccc Pro	cga Arg	aag Lys	acc Thr 230	cac His	tta Leu	ctg Leu	tac Tyr	atc Ile 235	ctc Leu	782
		tct Ser				tag	gggt	ggg	gacc	gggg	ag c	acct	gcct	g		830

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<211> 242

<212> PRT

<213> Homo sapiens

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Leu Leu Leu Leu Ser Thr Leu Val Ile Pro Ser Ala Ala Pro 20 25 30

Ile His Asp Ala Asp Ala Gln Glu Ser Ser Leu Gly Leu Thr Gly Leu 35 40 45

Gln Ser Leu Leu Gln Gly Phe Ser Arg Leu Phe Leu Lys Gly Asn Leu 50 55 60

Leu Arg Gly Ile Asp Ser Leu Phe Ser Ala Pro Met Asp Phe Arg Gly 65 70 75 80

Leu Pro Gly Asn Tyr His Lys Glu Glu Asn Gln Glu His Gln Leu Gly 85 90 95

Asn Asn Thr Leu Ser Ser His Leu Gln Ile Asp Lys Met Thr Asp Asn 100 105 110

Lys Thr Gly Glu Val Leu Ile Ser Glu Asn Val Val Ala Ser Ile Gln 115 120 125

Pro Ala Glu Gly Ser Phe Glu Gly Asp Leu Lys Val Pro Arg Met Glu 130 135 140

Glu Lys Glu Ala Leu Val Pro Ile Gln Lys Ala Thr Asp Ser Phe His 145 150 155 160

Thr Glu Leu His Pro Arg Val Ala Phe Trp Ile Ile Lys Leu Pro Arg 165 170 175

Arg Arg Ser His Gln Asp Ala Leu Glu Gly Gly His Trp Leu Ser Glu 180 185 190

Lys Arg His Arg Leu Gln Ala Ile Arg Asp Gly Leu Arg Lys Gly Thr 195 200 205

His Lys Asp Val Leu Glu Glu Gly Thr Glu Ser Ser His Ser Arg 210 215 220

Leu Ser Pro Arg Lys Thr His Leu Leu Tyr Ile Leu Arg Pro Ser Arg 225 230 235 240

Gln Leu

<210> 15

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200

195

cac aag gac gtc cta gaa gag ggg acc gag agc tcc tcc cac tcc agg His Lys Asp Val Leu Glu Glu Gly Thr Glu Ser Ser His Ser Arg 210 215 220	672
ctg tcc ccc cga aag acc cac tta ctg tac atc ctc agg ccc tct cgg Leu Ser Pro Arg Lys Thr His Leu Leu Tyr Ile Leu Arg Pro Ser Arg 225 230 235 240	720
cag ctg Gln Leu	726
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ctc ggg ggt att ttg ctg tgt aca ctg ctg gcg gcg gcg gtc ccc act Leu Gly Gly Ile Leu Leu Cys Thr Leu Leu Ala Ala Ala Val Pro Thr 5 10 15	165
gct cct gct cct tcc ccg acg gtc act tgg act ccg gcg gag ccg ggc Ala Pro Ala Pro Ser Pro Thr Val Thr Trp Thr Pro Ala Glu Pro Gly 20 25 30 35	213
cca gct ctc aac tac cct cag gag gaa gct acg ctc aat gag atg ttt Pro Ala Leu Asn Tyr Pro Gln Glu Glu Ala Thr Leu Asn Glu Met Phe 40 45 50	261
cga gag gtg gag ctg atg gaa gac act cag cac aaa ctg cgc agt Arg Glu Val Glu Glu Leu Met Glu Asp Thr Gln His Lys Leu Arg Ser 55 60 65	309
gcc gtg gag gag atg gag gcg gaa gaa gca gct gct aaa acg tcc tct Ala Val Glu Glu Met Glu Ala Glu Glu Ala Ala Ala Lys Thr Ser Ser 70 75 80	357
gag gtg aac ctg gca agc tta cct ccc aac tat cac aat gag acc agc Glu Val Asn Leu Ala Ser Leu Pro Pro Asn Tyr His Asn Glu Thr Ser 85 90 95	405
acg gag acc agg gtg gga aat aac aca gtc cat gtg cac cag gaa gtt Thr Glu Thr Arg Val Gly Asn Asn Thr Val His Val His Gln Glu Val 100 105 110 115	453
cac aag ata acc aac aac cag agt gga cag gtg gtc ttt tct gag aca His Lys Ile Thr Asn Asn Gln Ser Gly Gln Val Val Phe Ser Glu Thr 120 125 130	501

gtc Val	att Ile	aca Thr	tct Ser 135	gta Val	ggg	gat Asp	gaa Glu	gaa Glu 140	ggc Gly	aag Lys	agg Arg	agc Ser	cat His 145	gaa Glu	- 5 -	549
atc Ile	att Ile	gat Asp 150	gaa Glu	gac Asp	tgt Cys	ggg Gly	ccc Pro 155	acc Thr	agg Arg	tac Tyr	tgc Cys	cag Gln 160	ttc Phe	tcc Ser	agc Ser	597
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<211> 349

<212> PRT

<213> Homo sapiens

<400> 17

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Glu Pro Gly Pro Ala Leu Asn Tyr Pro Gln Glu Glu Ala Thr Leu Asn 35 40 45

Glu Met Phe Arg Glu Val Glu Glu Leu Met Glu Asp Thr Gln His Lys
50 55 60

Leu Arg Ser Ala Val Glu Glu Met Glu Ala Glu Glu Ala Ala Ala Lys 65 70 75 80

Thr Ser Ser Glu Val Asn Leu Ala Ser Leu Pro Pro Asn Tyr His Asn 85 90 95

Glu Thr Ser Thr Glu Thr Arg Val Gly Asn Asn Thr Val His Val His
100 105 110

Gln Glu Val His Lys Ile Thr Asn Asn Gln Ser Gly Gln Val Val Phe 115 120 125

Ser Glu Thr Val Ile Thr Ser Val Gly Asp Glu Glu Gly Lys Arg Ser 130 135 140

His Glu Cys Ile Ile Asp Glu Asp Cys Gly Pro Thr Arg Tyr Cys Gln 145 150 155 160

Phe Ser Ser Phe Lys Tyr Thr Cys Gln Pro Cys Arg Asp Gln Gln Met 165 170 175

Leu Cys Thr Arg Asp Ser Glu Cys Cys Gly Asp Gln Leu Cys Ala Trp 180 185 190

Gly His Cys Thr Gln Lys Ala Thr Lys Gly Gly Asn Gly Thr Ile Cys 195 200 205

Asp Asn Gln Arg Asp Cys Gln Pro Gly Leu Cys Cys Ala Phe Gln Arg 210 215 220

Gly Leu Leu Phe Pro Val Cys Thr Pro Leu Pro Val Glu Gly Glu Leu 225 230 235 240

Cys His Asp Pro Thr Ser Gln Leu Leu Asp Leu Ile Thr Trp Glu Leu 245 250 255

Glu Pro Glu Gly Ala Leu Asp Arg Cys Pro Cys Ala Ser Gly Leu Leu 260 265 270 .

Cys Gln Pro His Ser His Ser Leu Val Tyr Met Cys Lys Pro Ala Phe 275 280 285

Val Gly Ser His Asp His Ser Glu Glu Ser Gln Leu Pro Arg Glu Ala 290 295 300

Pro Asp Glu Tyr Glu Asp Val Gly Phe Ile Gly Glu Val Arg Gln Glu 305 310 315 320

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gtc cc Val Pr	c act o Thr	gct Ala 20	cct Pro	gct Ala	cct Pro	tcc Ser	ccg Pro 25	acg Thr	gtc Val	act Thr	tgg Trp	act Thr 30	ccg Pro	gcg Ala	96
gag cc Glu Pr	g ggc o Gly 35	Pro	gct Ala	ctc Leu	aac Asn	tac Tyr 40	cct Pro	cag Gln	gag Glu	gaa Glu	gct Ala 45	acg Thr	ctc Leu	aat Asn	144
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tct ga Ser Gl 13	u Thr	gtc Val	att Ile	aca Thr	tct Ser 135	gta Val	ggg	gat Asp	gaa Glu	gaa Glu 140	ggc Gly	aag Lys	agg Arg	agc Ser	432
cat ga His Gl 145	a tgt u Cys	atc Ile	att Ile	gat Asp 150	gaa Glu	gac Asp	tgt Cys	Gly	ccc Pro 155	acc Thr	agg Arg	tac Tyr	tgc Cys	cag Gln 160	480
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cta to Leu Cy	ge acc vs Thr	c cga Arg 180	gac Asp	agt Ser	gag Glu	tgc Cys	tgt Cys 185	gga Gly	gac Asp	cag Gln	ctg Leu	tgt Cys 190	gcc Ala	tgg Trp	576
ggt ca Gly Hi	s Cys	Thr	caa Gln	aag Lys	gcc Ala	acc Thr 200	aaa Lys	ggt Gly	ggc Gly	aat Asn	ggg Gly 205	acc Thr	atc Ile	tgt Cys	624
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ggc ct Gly Le 225	ig cto eu Lei	g ttc ı Phe	ccc Pro	gtg Val 230	tgc Cys	aca Thr	ccc Pro	ctg Leu	ccc Pro 235	gtg Val	gag Glu	gga Gly	gag Glu	ctc Leu 240	720

tgc cat gac ccc acc agc cag ctg ctg Cys His Asp Pro Thr Ser Gln Leu Leu 245	gat ctc atc acc tgg gaa ctg 768 Asp Leu Ile Thr Trp Glu Leu 250 255										
gag cct gaa gga gct ttg gac cga tgc Glu Pro Glu Gly Ala Leu Asp Arg Cys 260 265	Pro Cys Ala Ser Gly Leu Leu										
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gtg ggc agc cat gac cac agt gag gag Val Gly Ser His Asp His Ser Glu Glu 290 295	agc cag ctg ccc agg gag gcc 912 Ser Gln Leu Pro Arg Glu Ala 300										
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gcgggaccaa ggaggtgcgg ggcaagaagg aacggaagcg gtgcgatcca cagggctggg 420 ttttcttgca ccttgggtca cgcctccttg gcgagaaagc gcctcgcatt tgattgcttc 480 cagttattgc agaacttcct gtcctggtgg agaagcgggt ctcgcttggg ttccgctaat 540 ttctgtcctg aggcgtgaga ctgagttcat agggtcctgg gtccccgaac caggaagggt 600 tgagggaaca caatctgcaa gccccgcga cccaagtgag gggccccgtg ttggggtcct 660 coctcocttt gcattoccac coctcogggo tttgcgtctt cctggggacc c cct cgc Pro Arg 1 cgg gag atg gcc gcg ttg atg cgg agc aag gat tcg tcc tgc tgc ctg 765 Arg Glu Met Ala Ala Leu Met Arg Ser Lys Asp Ser Ser Cys Cys Leu ctc cta ctg gcc gcg gtg ctg atg gtg gag agc tca cag atc ggc agt 813 Leu Leu Leu Ala Ala Val Leu Met Val Glu Ser Ser Gln Ile Gly Ser tcg cgg gcc aaa ctc aac tcc atc aag tcc tct ctg ggc ggg gag acg 861 Ser Arg Ala Lys Leu Asn Ser Ile Lys Ser Ser Leu Gly Gly Glu Thr 40 cct ggt cag gcc gcc aat cga tct gcg ggc atg tac caa gga ctg gca 909 Pro Gly Gln Ala Ala Asn Arg Ser Ala Gly Met Tyr Gln Gly Leu Ala ttc ggc ggc agt aag aag ggc aaa aac ctg ggg cag gcc tac cct tgt 957 Phe Gly Gly Ser Lys Lys Gly Lys Asn Leu Gly Gln Ala Tyr Pro Cys agc agt gat aag gag tgt gaa gtt ggg agg tat tgc cac agt ccc cac 1005 Ser Ser Asp Lys Glu Cys Glu Val Gly Arg Tyr Cys His Ser Pro His 1053 caa gga tca tcg gcc tgc atg gtg tgt cgg aga aaa aag aag cgc tgc Gln Gly Ser Ser Ala Cys Met Val Cys Arg Arg Lys Lys Lys Arg Cys 105 cac cga gat ggc atg tgc tgc ccc agt acc cgc tgc aat aat ggc atc 1101 His Arg Asp Gly Met Cys Cys Pro Ser Thr Arg Cys Asn Asn Gly Ile 120 tgt atc cca gtt act gaa agc atc tta acc cct cac atc ccg gct ctg 1149 Cys Ile Pro Val Thr Glu Ser Ile Leu Thr Pro His Ile Pro Ala Leu 140 gat ggt act cgg cac aga gat cga aac cac ggt cat tac tca aac cat 1197 Asp Gly Thr Arg His Arg Asp Arg Asn His Gly His Tyr Ser Asn His gac ttg gga tgg cag aat cta gga aga cca cac act aag atg tca cat 1245 Asp Leu Gly Trp Gln Asn Leu Gly Arg Pro His Thr Lys Met Ser His 170 1293 ata aaa ggg cat gaa gga gac ccc tgc cta cga tca tca gac tgc att Ile Lys Gly His Glu Gly Asp Pro Cys Leu Arg Ser Ser Asp Cys Ile 185 180

gaa ggg ttt Glu Gly Phe 195	e Cys Cys A	ct cgt cat la Arg His 00	ttc tgg acc Phe Trp Thr 205	aaa atc tgc Lys Ile Cys	aaa cca Lys Pro 210	1341
gtg ctc cat Val Leu His	c cag ggg ga s Gln Gly G. 215	aa gtc tgt lu Val Cys	acc aaa caa Thr Lys Gln 220	cgc aag aag Arg Lys Lys	g ggt tct Gly Ser 225	1389
cat ggg cto His Gly Lev	g gaa att t 1 Glu Ile P 230	ne Gln Arg	tgc gac tgt Cys Asp Cys 235	gcg aag ggc Ala Lys Gly 240	Leu Ser	1437
tgc aaa gta Cys Lys Val 245	l Trp Lys A	at gcc acc sp Ala Thr 250	tac tcc tcc Tyr Ser Ser	aaa gcc aga Lys Ala Arc 255	a ctc cat g Leu His	1485
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<212> PRT

<213> Homo sapiens

<400> 21

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Gly Ser Ser Arg Ala Lys Leu Asn Ser Ile Lys Ser Ser Leu Gly Gly 35 40 45

Glu Thr Pro Gly Gln Ala Ala Asn Arg Ser Ala Gly Met Tyr Gln Gly
50 55 60

Leu Ala Phe Gly Gly Ser Lys Lys Gly Lys Asn Leu Gly Gln Ala Tyr 65 70 75 80

Pro Cys Ser Ser Asp Lys Glu Cys Glu Val Gly Arg Tyr Cys His Ser 85 90 95

Pro His Gln Gly Ser Ser Ala Cys Met Val Cys Arg Arg Lys Lys 100 105 110 Arg Cys His Arg Asp Gly Met Cys Cys Pro Ser Thr Arg Cys Asn Asn Gly Ile Cys Ile Pro Val Thr Glu Ser Ile Leu Thr Pro His Ile Pro 140 135 Ala Leu Asp Gly Thr Arg His Arg Asp Arg Asn His Gly His Tyr Ser Asn His Asp Leu Gly Trp Gln Asn Leu Gly Arg Pro His Thr Lys Met 170 Ser His Ile Lys Gly His Glu Gly Asp Pro Cys Leu Arg Ser Ser Asp 185 Cys Ile Glu Gly Phe Cys Cys Ala Arg His Phe Trp Thr Lys Ile Cys Lys Pro Val Leu His Gln Gly Glu Val Cys Thr Lys Gln Arg Lys Lys 215 Gly Ser His Gly Leu Glu Ile Phe Gln Arg Cys Asp Cys Ala Lys Gly 230 Leu Ser Cys Lys Val Trp Lys Asp Ala Thr Tyr Ser Ser Lys Ala Arg 250 245 Leu His Val Cys Gln Lys Ile 260 <210> 22 <211> 789 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (1)..(789) <400> 22 48 cct cgc cgg gag atg gcc gcg ttg atg cgg agc aag gat tcg tcc tgc Pro Arg Arg Glu Met Ala Ala Leu Met Arg Ser Lys Asp Ser Ser Cys 10 tgc ctg ctc cta ctg gcc gcg gtg ctg atg gtg gag agc tca cag atc 96 Cys Leu Leu Leu Ala Ala Val Leu Met Val Glu Ser Ser Gln Ile 20 25 30 ggc agt tcg cgg gcc aaa ctc aac tcc atc aag tcc tct ctg ggc ggg 144 Gly Ser Ser Arg Ala Lys Leu Asn Ser Ile Lys Ser Ser Leu Gly Gly 35 gag acg cct ggt cag gcc gcc aat cga tct gcg ggc atg tac caa gga 192 Glu Thr Pro Gly Gln Ala Ala Asn Arg Ser Ala Gly Met Tyr Gln Gly 55 50 240 ctg gca ttc ggc ggc agt aag aag ggc aaa aac ctg ggg cag gcc tac Leu Ala Phe Gly Gly Ser Lys Lys Gly Lys Asn Leu Gly Gln Ala Tyr 70 65

cct tgt agc Pro Cys Ser	agt gat aag Ser Asp Lys 85	gag tgt gaa Glu Cys Glu	gtt ggg ag Val Gly An 90	gg tat tgc rg Tyr Cys	cac agt 28 His Ser 95	38
ccc cac caa Pro His Gln	gga tca tcg Gly Ser Ser 100	gcc tgc atg Ala Cys Met 105	gtg tgt co Val Cys Ai	gg aga aaa rg Arg Lys 110	aag aag 33 Lys Lys	36
cgc tgc cac Arg Cys His 115	cga gat ggc Arg Asp Gly	atg tgc tgc Met Cys Cys 120	ccc agt ac Pro Ser Th	cc cgc tgc hr Arg Cys 125	aat aat 38 Asn Asn	34
ggc atc tgt Gly Ile Cys · 130	atc cca gtt Ile Pro Val	act gaa agc Thr Glu Ser 135	Ile Leu Th	cc cct cac hr Pro His 40	atc ccg 43 Ile Pro	32
gct ctg gat Ala Leu Asp 145	ggt act cgg Gly Thr Arg 150	cac aga gat His Arg Asp	cga aac ca Arg Asn H: 155	ac ggt cat is Gly His	tac tca 48 Tyr Ser 160	30
aac cat gac Asn His Asp	ttg gga tgg Leu Gly Trp 165	cag aat cta Gln Asn Leu	gga aga co Gly Arg P: 170	ca cac act ro His Thr	aag atg 52 Lys Met 175	28
tca cat ata Ser His Ile	aaa ggg cat Lys Gly His 180	gaa gga gac Glu Gly Asp 185	Pro Cys L	ta cga tca eu Arg Ser 190	tca gac 5° Ser Asp	76
tgc att gaa Cys Ile Glu 195	ggg ttt tgc Gly Phe Cys	tgt gct cgt Cys Ala Arg 200	cat ttc to His Phe T	gg acc aaa rp Thr Lys 205	atc tgc 62 Ile Cys	24
aaa cca gtg Lys Pro Val 210	ctc cat cag Leu His Gln	ggg gaa gtc Gly Glu Val 215	Cys Thr L	aa caa cgc ys Gln Arg 20	aag aag 6 Lys Lys	72
ggt tct cat Gly Ser His 225	ggg ctg gaa Gly Leu Glu 230	Ile Phe Gln	cgt tgc g Arg Cys A 235	ac tgt gcg .sp Cys Ala	aag ggc 72 Lys Gly 240	20
ctg tct tgc Leu Ser Cys	aaa gta tgg Lys Val Trp 245	aaa gat gcc Lys Asp Ala	acc tac to Thr Tyr So 250	cc tcc aaa er Ser Lys	_	68
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<211> 54

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: consensus sequence

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<223> Xaa's at positions 2,3,5,6,8-12,14-26,28,29,31-36,38-42,45-48, and

50-53 may be any amino acid

<220>

<223> Xaa's at postions 22-26 may be absent

<400> 23

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Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Cys Xaa Cys Xaa Xaa Cys Xaa Xaa 20 25 30

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Cys Xaa Xaa Xaa Cys 50

<210> 24

<211> 123

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: consensus
 sequence

<220>

<223> Xaa's at positions 2,5,8,9,11,14-23,25,27-30,32,33,35-53,60,62,63,65, 68,70,71,73-96,98,100,101,104, and 106-122 may be any amino acid

<220>

<223> Xaa's at positions 22,23,51-58,89-96 and 116-122 may be absent

<400> 24

Cys Xaa Xaa Xaa Xaa Asp Cys Xaa Xaa Gly Xaa Cys Cys Xaa Xaa Xaa 1 5 10 15

Xaa Xaa Xaa Xaa Xaa Xaa Cys Xaa Pro Xaa Xaa Xaa Gly Xaa 20 25 30

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Cys Xaa Cys Xaa Xaa Pro 50 55 60 .

Cys Xaa Cys Xaa Xaa Gly Leu Xaa Cys Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa 100 105 110

35

50

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cta aaa aat gac ctg cta cga gac ctg gac aac ttc ttc tcc tcc ccc

Leu Lys Asn Asp Leu Leu Arg Asp Leu Asp Asn Phe Phe Ser Ser Pro

251

atg gac ttc cg. Met Asp Phe Arc											299
gag cac aga at Glu His Arg Me 8	Gly Asn										347
aag gtg act ga Lys Val Thr As 100	c aac cag o Asn Gln	aca ggg Thr Gly 105	gag Glu	gtg Val	cac His	atc Ile	tcg Ser 110	gag Glu	aaa Lys	gtc Val	395
gag gcc tcc at Glu Ala Ser Il 115											443
ccc aaa gta ga Pro Lys Val Gl 130	a gca aaa 1 Ala Lys 135	gag ccc Glu Pro	ccg Pro	gtg Val	cct Pro 140	gtg Val	cag Gln	aag Lys	gtc Val	acc Thr 145	491
gac agc ttg ca Asp Ser Leu Hi											539
atg cca agg cg Met Pro Arg Ar 16	g Arg Thr	cag ccc Gln Pro	gat Asp 170	gtc Val	cag Gln	gat Asp	gga Gly	ggc Gly 175	cgc Arg	tgg Trp	587
ctc ata gaa aa Leu Ile Glu Ly 180											635
gga ggc gcc cg Gly Gly Ala Ar 195											683
cac gcc aag ct His Ala Lys Le 210	g cct gtc u Pro Val 215	aga aag Arg Lys	aca Thr	cac His	ttt Phe 220	ctc Leu	tac Tyr	atc Ile	ctc Leu	agg Arg 225	731
cca tcc caa ca Pro Ser Gln Gl		gtgggga	ccaga	atgto	cc ca	acac	ccta	C CC	caaca	acca	786
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Ser Ser Ser Al	_	Ile His	Asp 25	Val	Asp	Ser	Gln	Gln 30	Asn	Thr	
Ser Gly Phe Le	u Gly Leu	Gln Arg 40		Leu	Gln	Ser	Phe 45	Ser	Arg	Leu	

i

Phe Leu Lys Asn Asp Leu Leu Arg Asp Leu Asp Asn Phe Phe Ser Ser Pro Met Asp Phe Arg Asp Leu Pro Arg Asn Phe His Gln Glu Glu Asn Gln Glu His Arg Met Gly Asn His Thr Leu Ser Ser His Leu Gln Ile Asp Lys Val Thr Asp Asn Gln Thr Gly Glu Val His Ile Ser Glu Lys 105 Val Glu Ala Ser Ile Glu Pro Glu Arg Asn Pro Glu Gly Asp Trp Lys Val Pro Lys Val Glu Ala Lys Glu Pro Pro Val Pro Val Gln Lys Val 135 Thr Asp Ser Leu His Pro Glu Pro Arg Gln Val Ala Phe Trp Ile Met 150 Lys Met Pro Arg Arg Arg Thr Gln Pro Asp Val Gln Asp Gly Gly Arg 170 165 Trp Leu Ile Glu Lys Arg His Arg Met Gln Ala Ile Arg Asp Gly Leu 180 185 Arg Gly Gly Ala Arg Glu Asp Ser Leu Glu Asp Gly Val His Ile Pro 200 195 Gln His Ala Lys Leu Pro Val Arg Lys Thr His Phe Leu Tyr Ile Leu 220 215 Arg Pro Ser Gln Gln Leu 225 230 <210> 28 <211> 690 <212> DNA <213> Mus musculus <220> <221> CDS <222> (1)..(690) <400> 28 atg tgt cga ctg agg gtc ttg ctg ctg ctc ccc ttg gcc ttc gtg 48 Met Cys Arg Leu Arg Val Leu Leu Leu Leu Pro Leu Ala Phe Val 10 tcc tcc tct gct ctc ccc atc cat gat gtc gac tct cag cag aac acc 96 Ser Ser Ser Ala Leu Pro Ile His Asp Val Asp Ser Gln Gln Asn Thr 25 tcc ggg ttc ctg ggc ctt cag agg ctt ctc caa agc ttt agt cga ctg 144 Ser Gly Phe Leu Gly Leu Gln Arg Leu Leu Gln Ser Phe Ser Arg Leu 40 35

ttc c Phe I	cta Leu 50	aaa Lys	aat Asn	gac Asp	ctg Leu	cta Leu 55	cga Arg	gac Asp	ctg Leu	gac Asp	aac Asn 60	ttc Phe	ttc Phe	tcc Ser	tcc Ser	192
ccc a Pro M 65	atg Met	gac Asp	ttc Phe	cga Arg	gac Asp 70	ctt Leu	cct Pro	agg Arg	aac Asn	ttc Phe 75	cat His	cag Gln	gaa Glu	gag Glu	aac Asn 80	240
cag g Gln G	gag Glu	cac His	aga Arg	atg Met 85	ggc Gly	aac Asn	cat His	acc Thr	ctc Leu 90	tcc Ser	agc Ser	cac His	cta Leu	cag Gln 95	ata Ile	288
gac a Asp I	aag Lys	gtg Val	act Thr 100	gac Asp	aac Asn	cag Gln	aca Thr	ggg Gly 105	gag Glu	gtg Val	cac His	atc Ile	tcg Ser 110	gag Glu	aaa Lys	336
gtc g Val G																384
gtt c Val E	ccc Pro 130	aaa Lys	gta Val	gaa Glu	gca Ala	aaa Lys 135	gag Glu	ccc Pro	ccg Pro	gtg Val	cct Pro 140	gtg Val	cag Gln	aag Lys	gtc Val	432
acc o Thr <i>H</i> 145																480
aag a Lys N	atg Met	cca Pro	agg Arg	cgg Arg 165	agg Arg	acc Thr	cag Gln	ccc Pro	gat Asp 170	gtc Val	cag Gln	gat Asp	gga Gly	ggc Gly 175	cgc Arg	528
tgg d Trp I	ctc Leu	ata Ile	gaa Glu 180	aag Lys	cga Arg	cat His	cgc Arg	atg Met 185	cag Gln	gcc Ala	atc Ile	cgg Arg	gat Asp 190	ggg	ctc Leu	576
cgt o	gga Gly	ggc Gly 195	gcc Ala	cgt Arg	gag Glu	gac Asp	agc Ser 200	ctg Leu	gag Glu	gat Asp	ggg Gly	gtc Val 205	cat His	atc Ile	ccc Pro	624
caa d Gln H	cac His 210	gcc Ala	aag Lys	ctg Leu	cct Pro	gtc Val 215	aga Arg	aag Lys	aca Thr	cac His	ttt Phe 220	, ctc Leu	tac Tyr	atc Ile	ctc Leu	· 672
agg d Arg I 225																690
<210> 29 <211> 51 <212> PRT <213> Artificial Sequence																
<220> <223> Xaa at positions 3-5,7,9-15,18,20-22,24-27,29,31,33,34,36-39,42,44, 45, and 47-50 may be any amino acid																

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<220>
<223> Description of Artificial Sequence: consensus
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Asn Xaa Thr Xaa Xaa Xaa His Xaa Xaa Xaa Lys Xaa Thr Xaa Asn
Xaa Xaa Gly Xaa Xaa Xaa Ser Glu Xaa Val Xaa Xaa Ser Xaa Xaa
Xaa Xaa Glu
     50
<210> 30
<211> 20
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: primer
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                                                                   20
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<210> 31
<211> 20
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<213> Artificial Sequence
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<400> 31
                                                                   20
tcttcagtca ggctcctctc
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<211> 21
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: primer
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                                                                   21
acctgcaatg tgtcgactga g
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20

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Asp His Ser His Phe Pro Arg Gly Glu Ile Glu Glu Ser Ile Ile Glu 145 150 155 160

Asn Leu Gly Asn Asp His Asn Ala Ala Ala Gly Asp Gly Tyr Pro Arg 165 170 175

Arg Thr Thr Leu Thr Ser Lys Ile Tyr His Thr Lys Gly Gln Glu Gly 180 185 190

Ser Val Cys Leu Arg Ser Ser Asp Cys Ala Ala Gly Leu Cys Cys Ala 195 200 205

Arg His Phe Trp Ser Lys Ile Cys Lys Pro Val Leu Lys Glu Gly Gln 210 215 220

Val Cys Thr Lys His Lys Arg Lys Gly Ser His Gly Leu Glu Ile Phe 225 230 235 240

Gln Arg Cys Tyr Cys Gly Glu Gly Leu Ala Cys Arg Ile Gln Lys Asp 245 250 255

His His Gln Ala Ser Asn Ser Ser Arg Leu His Thr Cys Gln Arg His 260 265 270

<210> 37

<211> 259

<212> PRT

<213> Xenopus laevis

<400> 37

Met Gly Ser Asn Met Phe Pro Val Pro Leu Ile Val Phe Trp Gly Phe 1 5 10 15

Ile Leu Asp Gly Ala Leu Gly Phe Val Met Met Thr Asn Ser Asn Ser 20 25 30

Ile Lys Asn Val Pro Ala Ala Pro Ala Gly Gln Pro Ile Gly Tyr Tyr
35 40 45

Pro Val Ser Val Ser Pro Asp Ser Leu Tyr Asp Ile Ala Asn Lys Tyr 50 55 60

Gln Pro Leu Asp Ala Tyr Pro Leu Tyr Ser Cys Thr Glu Asp Asp Asp 65 70 75 80

Cys Ala Leu Asp Glu Phe Cys His Ser Ser Arg Asn Gly Asn Ser Leu 85 90 95

Val Cys Leu Ala Cys Arg Lys Arg Lys Arg Cys Leu Arg Asp Ala 100 105 110

Met Cys Cys Thr Gly Asn Tyr Cys Ser Asn Gly Ile Cys Val Pro Val 115 120 125

Glu Gln Asp Gln Glu Arg Phe Gln His Gln Gly Tyr Leu Glu Glu Thr 130 135 140

Ile Leu Glu Asn Tyr Asn Asn Ala Asp His Ala Thr Met Asp Thr His 145 150 155 160

Ser Lys Leu Thr Thr Ser Pro Ser Gly Met Gln Pro Phe Lys Gly Arg 165 170 175

Asp Gly Asp Val Cys Leu Arg Ser Thr Asp Cys Ala Pro Gly Leu Cys 180 185 190

Cys Ala Arg His Phe Trp Ser Lys Ile Cys Lys Pro Val Leu Asp Glu 195 200 205

Gly Gln Val Cys Thr Lys His Arg Arg Lys Gly Ser His Gly Leu Glu 210 215 220

Ile Phe Gln Arg Cys His Cys Gly Ala Gly Leu Ser Cys Arg Leu Gln 225 230 235 240

Lys Gly Glu Phe Thr Thr Val Pro Lys Thr Ser Arg Leu His Thr Cys $245 \hspace{1.5cm} 250 \hspace{1.5cm} 255$

Gln Arg His

<210> 38

<211> 350

<212> PRT

<213> Gallus gallus

<400> 38

Met Arg Arg Gly Glu Gly Pro Ala Pro Arg Arg Arg Trp Leu Leu 1 5 10 15

Leu Ala Val Leu Ala Ala Leu Cys Cys Ala Ala Ala Gly Ser Gly Gly 20 25 30

Arg Arg Ala Ala Ser Leu Gly Glu Met Leu Arg Glu Val Glu Ala 35 40 45

Leu Met Glu Asp Thr Gln His Lys Leu Arg Asn Ala Val Gln Glu Met 50 60

Glu Ala Glu Glu Glu Gly Ala Lys Lys Leu Ser Glu Val Asn Phe Glu 65 70 75 80

Asn Leu Pro Pro Thr Tyr His Asn Glu Ser Asn Thr Glu Thr Arg Ile 85 90 95

Gly Asn Lys Thr Val Gln Thr His Gln Glu Ile Asp Lys Val Thr Asp 100 105 110

Asn Arg Thr Gly Ser Thr Ile Phe Ser Glu Thr Ile Ile Thr Ser Ile 115 120 125

Lys Gly Gly Glu Asn Lys Arg Asn His Glu Cys Ile Ile Asp Glu Asp 130 135 140

Cys Glu Thr Gly Lys Tyr Cys Gln Phe Ser Thr Phe Glu Tyr Lys Cys 145 150 155 160

Gln Pro Cys Lys Thr Gln His Thr His Cys Ser Arg Asp Val Glu Cys 165 170 175 Cys Gly Asp Gln Leu Cys Val Trp Gly Glu Cys Arg Lys Ala Thr Ser Arg Gly Glu Asn Gly Thr Ile Cys Glu Asn Gln His Asp Cys Asn Pro 200 Gly Thr Cys Cys Ala Phe Gln Lys Glu Leu Leu Phe Pro Val Cys Thr Pro Leu Pro Glu Glu Gly Glu Pro Cys His Asp Pro Ser Asn Arg Leu Leu Asn Leu Ile Thr Trp Glu Leu Glu Pro Asp Gly Val Leu Glu Arg Cys Pro Cys Ala Ser Gly Leu Ile Cys Gln Pro Gln Ser Ser His Ser Thr Thr Ser Val Cys Glu Leu Ser Ser Asn Glu Thr Arg Lys Asn Glu 280 Lys Glu Asp Pro Leu Asn Met Asp Glu Met Pro Phe Ile Ser Leu Ile 295 Pro Arg Asp Ile Leu Ser Asp Tyr Glu Glu Ser Ser Val Ile Gln Glu 310 Val Arg Lys Glu Leu Glu Ser Leu Glu Asp Gln Ala Gly Val Lys Ser 330 325 Glu His Asp Pro Ala His Asp Leu Phe Leu Gly Asp Glu Ile 345 340

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